

10/519,931

FILE 'HOME' ENTERED AT 12:16:11 ON 10 OCT 2007

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SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

0.21

0.21

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DICTIONARY FILE UPDATES: 9 OCT 2007 HIGHEST RN 949922-95-6

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=>

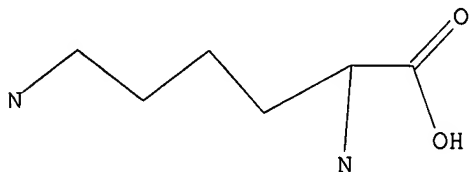
Uploading C:\Program Files\Stnexp\Queries\10519931.str

L1 STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1 STR



Structure attributes must be viewed using STN Express query preparation.

=> s l1 full

FULL SEARCH INITIATED 12:16:58 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 469999 TO ITERATE

100.0% PROCESSED 469999 ITERATIONS

83880 ANSWERS

SEARCH TIME: 00.00.06

L2 83880 SEA SSS FUL L1

file caplus  
COST IN U.S. DOLLARS

| SINCE FILE | TOTAL   |
|------------|---------|
| ENTRY      | SESSION |
| 172.10     | 172.31  |

FULL ESTIMATED COST

FILE 'CAPLUS' ENTERED AT 12:17:10 ON 10 OCT 2007  
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FILE COVERS 1907 - 10 Oct 2007 VOL 147 ISS 16  
FILE LAST UPDATED: 9 Oct 2007 (20071009/ED)

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<http://www.cas.org/infopolicy.html>

=> s 12

L3 107468 L2

=> s 13 and fmoc

6432 FMOC

L4 1286 L3 AND FMOC

=> s 14 and PNA

6518 PNA

L5 27 L4 AND PNA

=> dup rem 15

PROCESSING COMPLETED FOR L5

L6 27 DUP REM L5 (0 DUPLICATES REMOVED)

=> s 16 and 2003/py

L7 27 S L6

1265272 2003/PY

L8 7 L7 AND 2003/PY

=> d 18 bib abs hitstr 1-7

L8 ANSWER 1 OF 7 CAPLUS COPYRIGHT 2007 ACS on STN

AN 2003:829485 CAPLUS

DN 140:111672

TI Synthesis of Radiometal-Labeled and Fluorescent Cell-Permeating Peptide-PNA Conjugates for Targeting the bcl-2 Proto-oncogene

AU Gallazzi, Fabio; Wang, Yi; Jia, Fang; Shenoy, Nalini; Landon, Linda A.; Hannink, Mark; Lever, Susan Z.; Lewis, Michael R.

CS Molecular Biology Program, Department of Veterinary Medicine and Surgery, Department of Chemistry, Department of Biochemistry, University of Missouri-Columbia, Columbia, MO, 65211, USA

CO Bioconjugate Chemistry (2003), 14(6), 1083-1095

CODEN: BCCHES; ISSN: 1043-1802

PB American Chemical Society

DT Journal

LA English

OS CASREACT 140:111672

AB The B-cell lymphoma/leukemia-2 (bcl-2) proto-oncogene has been associated with the transformation of benign lesions to malignancy, disease progression, poor prognosis, reduced survival, and development of resistance to radiation and chemotherapy in many types of cancer. The objective of this work was to synthesize an antisense peptide nucleic acid (PNA) complementary to the first six codons of the bcl-2 open reading frame, conjugated to a membrane-permeating peptide for intracellular delivery, and modified with a bifunctional chelating agent for targeting imaging and therapeutic radiometals to tumors overexpressing bcl-2. Four peptide-PNA constructs were synthesized by a combination of manual and automated stepwise elongation techniques, including bcl-2 antisense conjugates and nonsense conjugates with no complementarity to any known mammalian gene or DNA sequence. The PNA sequences were synthesized manually by solid-phase 9-fluorenylmethoxycarbonyl (Fmoc) techniques. Then a fully protected lysine monomer, modified with 1,4,7,10-tetraazacyclododecane-N,N',N'',N'''-tetraacetic acid (DOTA) for radiometal chelation, was coupled manually to each PNA sequence. Synthesis of the DOTA-PNA conjugates was followed by automated elongation with a peptide sequence (PTD-4-glycine, PTD-4-G), known to mediate cellular internalization of impermeable effector mols., or its retro-inverso analog (ri-PTD-4-G). Preparation of the four conjugates required an innovative synthetic strategy, using mild acid conditions to generate hydrophobic, partially deprotected intermediates. These intermediates were purified by semipreparative reversed-phase HPLC and completely deprotected to yield pure peptide-PNA conjugates in 6% to 9% overall yield. Using modifications of this synthetic strategy, the ri-PTD-4-G conjugate of bcl-2 antisense PNA was prepared using a lysine derivative of tetramethylrhodamine (TMR) for fluorescence microscopy. Plasma stability studies showed that <sup>111</sup>In-DOTA-labeled ri-PTD-4-G-anti-bcl-2 PNA was stable for 168 h at 37 °C, unlike the conjugate containing the parent peptide sequence. Scanning confocal fluorescence microscopy of TMR-labeled ri-PTD-4-G-anti-bcl-2 PNA in Raji lymphoma cells demonstrated that the retro-inverso peptide was active in membrane permeation and mediated cellular internalization of the antisense PNA into the cytoplasm, where high concns. of bcl-2 mRNA are expected to be present.

IT 105047-45-8 635732-47-7

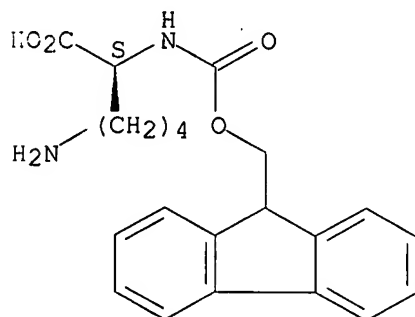
RL: RCT (Reactant); RACT (Reactant or reagent)

(synthesis of radiometal-labeled and fluorescent cell-permeating peptide-PNA conjugates for targeting the bcl-2 proto-oncogene)

RN 105047-45-8 CAPLUS

CN L-Lysine, N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (CA INDEX NAME)

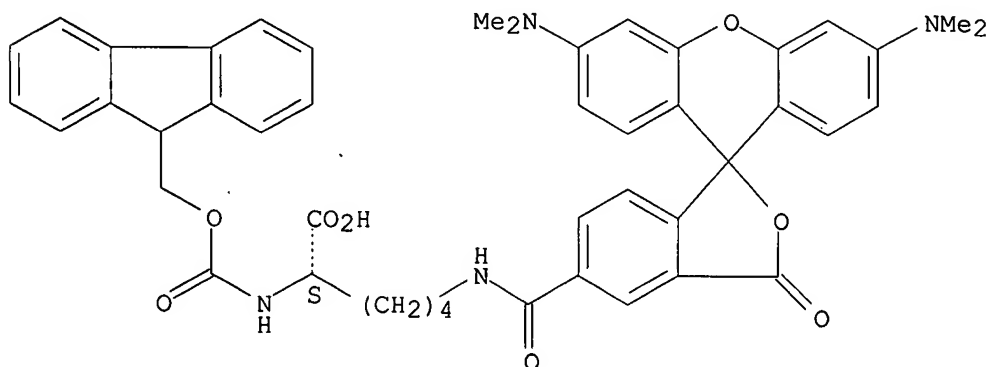
Absolute stereochemistry.



RN 635732-47-7 CAPLUS

CN L-Lysine, N6-[[3',6'-bis(dimethylamino)-3-oxospiro[isobenzofuran-1(3H),9'-[9H]xanthen]-5-yl]carbonyl]-N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (CA INDEX NAME)

Absolute stereochemistry.



635732-44-4P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(synthesis of radiometal-labeled and fluorescent cell-permeating peptide-PNA conjugates for targeting the bcl-2 proto-oncogene)

RN 635732-44-4 CAPLUS

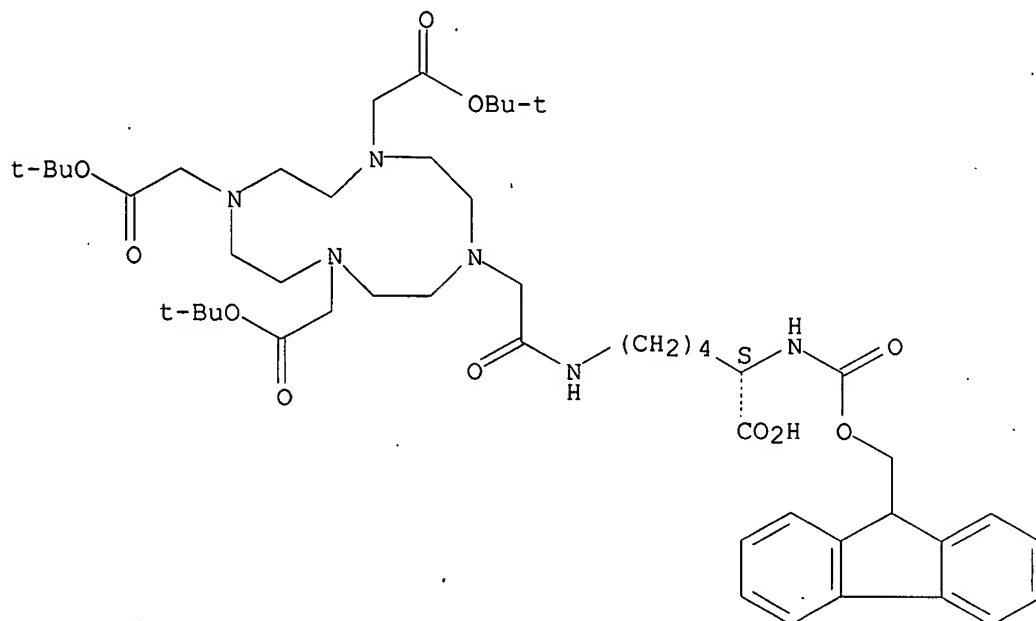
CN 1,4,7,10-Tetraazacyclododecane-1,4,7-triacetic acid, 10-[2-[[[(5S)-5-carboxy-5-[[[(9H-fluoren-9-ylmethoxy)carbonyl]amino]pentyl]amino]-2-oxoethyl]-,  $\alpha,\alpha',\alpha''$ -tris(1,1-dimethylethyl) ester, tetrakis(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 479081-06-6

CMF C49 H74 N6 O11

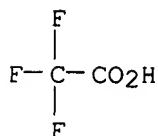
Absolute stereochemistry.



CM 2

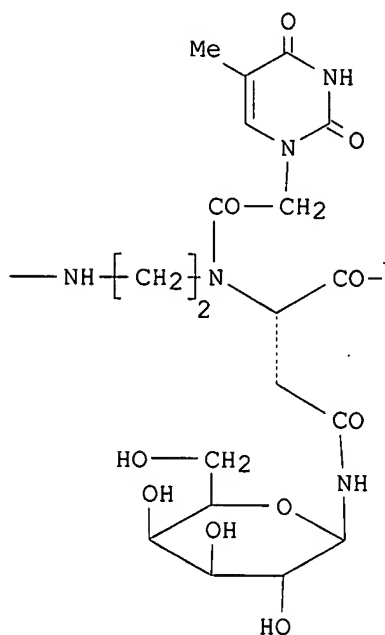
CRN 76-05-1

CMF C2 H F3 O2



..L.CNT 58 THERE ARE 58 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 2 OF 7 CAPLUS COPYRIGHT 2007 ACS on STN  
AN 2003:679388 CAPLUS  
DN 139:381726  
TI Modulation of the Pharmacokinetic Properties of PNA: Preparation  
of Galactosyl, Mannosyl, Fucosyl, N-Acetylgalactosaminy, and  
N-Acetylglucosaminy Derivatives of Aminoethylglycine Peptide Nucleic Acid  
Monomers and Their Incorporation into PNA Oligomers  
AU Hamzavi, Ramin; Dolle, Frederic; Tavitian, Bertrand; Dahl, Otto; Nielsen,  
Peter E.  
CS Center for Biomolecular Recognition, Department of Medical Biochemistry  
and Genetics, University of Copenhagen, Copenhagen, DK-2200, Den.  
SO Bioconjugate Chemistry (2003), 14(5), 941-954  
CODEN: BCCHES; ISSN: 1043-1802  
PB American Chemical Society  
DT Journal  
LA English  
CS CASREACT 139:381726  
GI



AB A series of N-(2-aminoethyl)- $\alpha$ -amino acid thymine peptide nucleic acid (PNA) monomers bearing glycosylated side chains in the

$\alpha$ -amino acid position (e.g, I) have been synthesized. These include PNA monomers where glycine has been replaced by serine and threonine (O-glycosylated), derivs. of lysine and nor-alanine (C-glycosylated), and amide derivs. of aspartic acid (N-glycosylated). The Boc and Fmoc derivs. of these monomers were used for incorporation in PNA oligomers. Twelve PNA decamers containing the glycosylated units in one, two, or three positions were prepared, and the thermal stability ( $T_m$ ) of their complexes with a complementary RNA was determined. Incorporation of the glycosyl monomers reduced the duplex stability by 0-6° C per substitution. A cysteine was attached to the amino terminus of eight of the PNA decamers (Cys-CTCATACTCT-NH<sub>2</sub>) for easy conjugation to a [<sup>18</sup>F]radiolabeled N-(4-fluorobenzyl)-2-bromoacetamide. The in vivo biodistribution of these PNA oligomers was determined in rat 2 h after i.v. administration. Most of the radioactivity was recovered in the kidneys and in the urine. However, N-acetylgalactosamine (and to a lesser extent galactose and mannose)-modified PNAs were effectively targeting the liver (40-fold over unmodified PNA). Thus, the pharmacodistribution in rats of PNA oligomers can be profoundly changed by glycosylation. These results could be of great significance for PNA drug development, as they should allow modulation and fine-tuning of the pharmacokinetic profile of a drug lead.

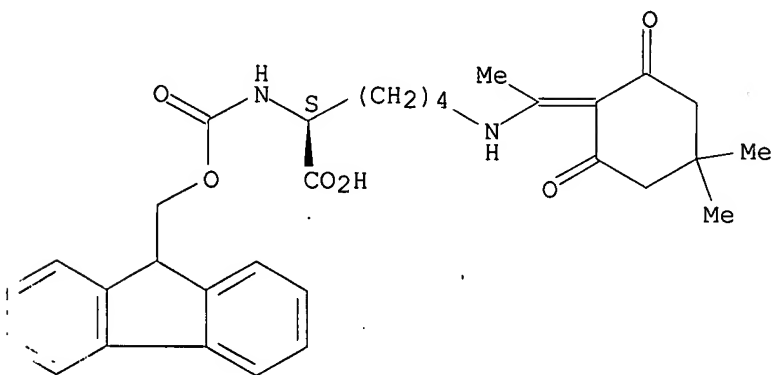
150629-67-7

RL: RCT (Reactant); RACT (Reactant or reagent)  
(preparation of glycosylated monomers for PNA synthesis and their effect on PNA/RNA hybridization or PNA biodistribution)

RN 150629-67-7 CAPLUS

CN L-Lysine, N6-[1-(4,4-dimethyl-2,6-dioxocyclohexylidene)ethyl]-N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (CA INDEX NAME)

Absolute stereochemistry.



IT 612491-20-0P 612491-21-1P 612491-22-2P

612491-23-3P 612491-24-4P 612491-25-5P

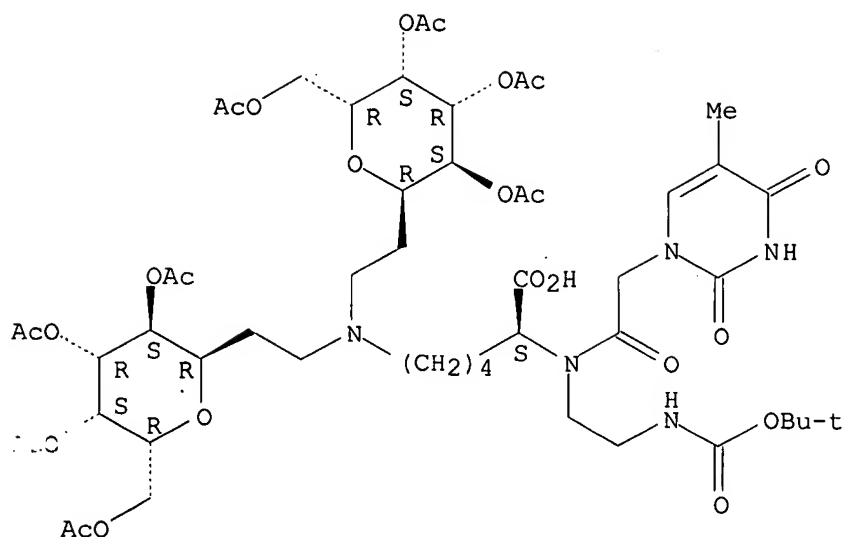
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of glycosylated monomers for PNA synthesis and their effect on PNA/RNA hybridization or PNA biodistribution)

RN 612491-20-0 CAPLUS

CN L-Lysine, N2-[(3,4-dihydro-5-methyl-2,4-dioxo-1(2H)-pyrimidinyl)acetyl]-N2-[2-[[[1,1-dimethylethoxy)carbonyl]amino]ethyl]-N6,N6-bis(1,3,4,5-tetra-O-acetyl-2,6-anhydro-7,8-dideoxy-D-glycero-L-galacto-octitol-8-yl)- (9CI)  
(CA INDEX NAME)

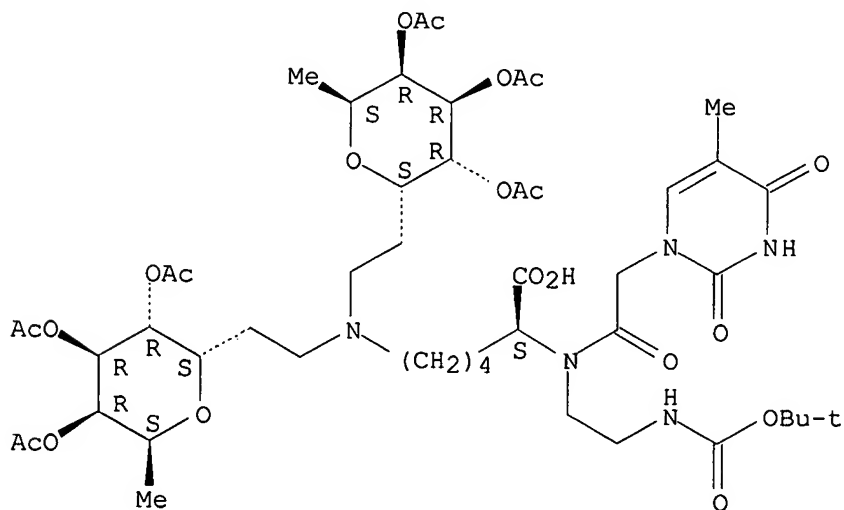
Absolute stereochemistry.



RN 612491-21-1 CAPLUS

CN L-Lysine, N2-[(3,4-dihydro-5-methyl-2,4-dioxo-1(2H)-pyrimidinyl)acetyl]-N2-[2-[[[(1,1-dimethylethoxy)carbonyl]amino]ethyl]-N6,N6-bis(3,4,5-tri-O-acetyl-2,6-anhydro-1,7,8-trideoxy-L-glycero-D-galacto-octitol-8-yl)- (9CI)  
(CA INDEX NAME)

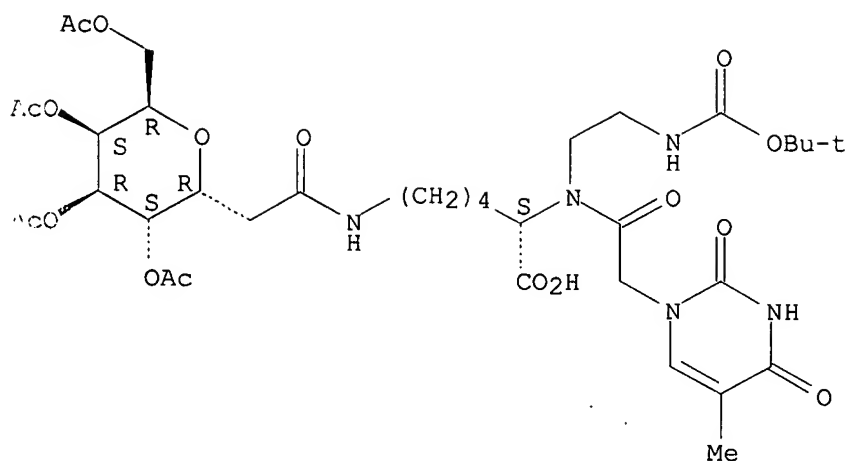
Absolute stereochemistry.



RN 612491-22-2 CAPLUS

CN L-Lysine, N2-[(3,4-dihydro-5-methyl-2,4-dioxo-1(2H)-pyrimidinyl)acetyl]-N2-[2-[[[(1,1-dimethylethoxy)carbonyl]amino]ethyl]-N6-(4,5,6,8-tetra-O-acetyl-3,7-anhydro-2-deoxy-D-glycero-L-glucopyranosyl)- (9CI) (CA INDEX NAME)

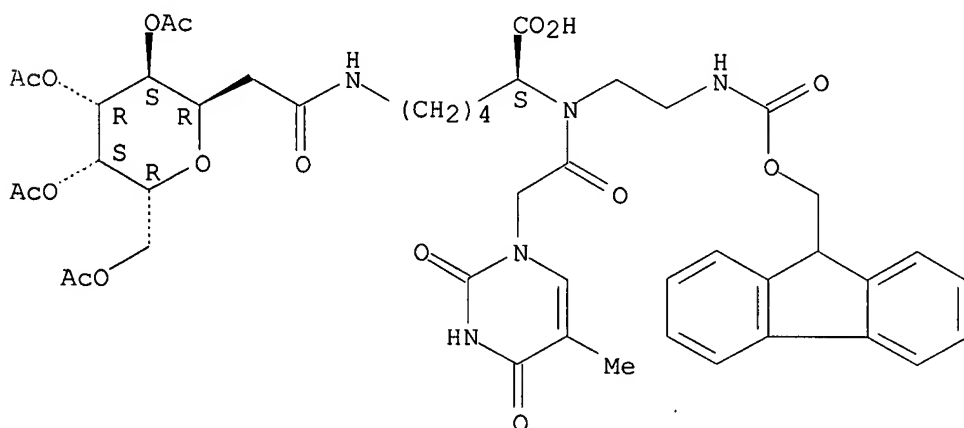
Absolute stereochemistry.



RN 612491-23-3 CAPLUS

CN L-Lysine, N2-[(3,4-dihydro-5-methyl-2,4-dioxo-1(2H)-pyrimidinyl)acetyl]-N2-[2-[[ (9H-fluoren-9-ylmethoxy) carbonyl] amino]ethyl]-N6-(4,5,6,8-tetra-O-acetyl-3,7-anhydro-2-deoxy-D-glycero-L-glucopyranosyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

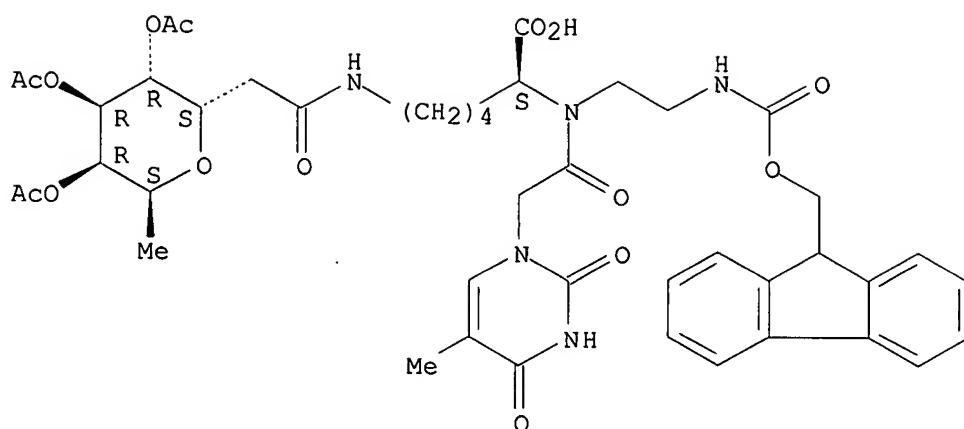


RN 612491-24-4 CAPLUS

CN L-Lysine, N2-[(3,4-dihydro-5-methyl-2,4-dioxo-1(2H)-pyrimidinyl)acetyl]-N2-[2-[[ (9H-fluoren-9-ylmethoxy) carbonyl] amino]ethyl]-N6-(4,5,6-tri-O-acetyl-3,7-anhydro-2,8-dideoxy-L-glycero-D-glucopyranosyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

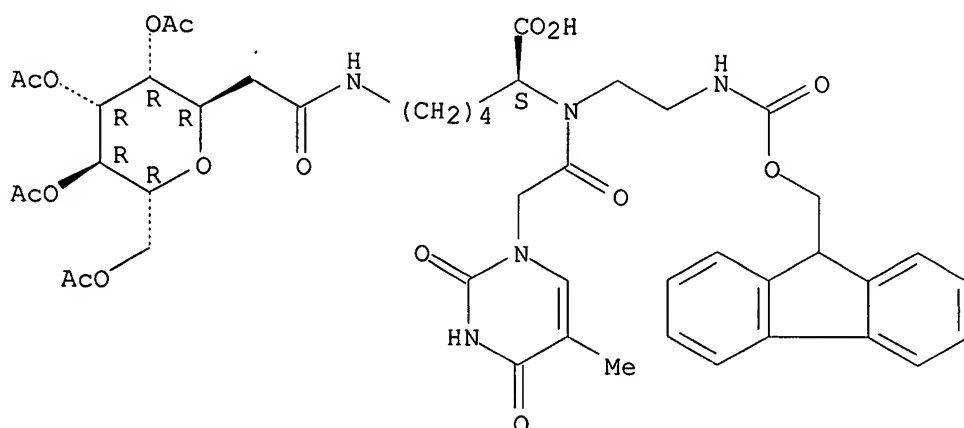




612491-25-5 CAPLUS

CN L-Lysine, N2-[(3,4-dihydro-5-methyl-2,4-dioxo-1(2H)-pyrimidinyl)acetyl]-N2-[2-[(9H-fluoren-9-ylmethoxy)carbonyl]amino]ethyl]-N6-(4,5,6,8-tetra-O-acetyl-3,7-anhydro-2-deoxy-D-glycero-D-talo-octonoyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RE.CNT 40 THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 3 OF 7 CAPLUS COPYRIGHT 2007 ACS on STN  
AN 2003:521324 CAPLUS  
DN 139:292469  
TI Synthesis and DNA binding properties of terminally modified peptide nucleic acids  
AU Mokhir, Andriy; Zohm, Burkhard; Fuessler, Andreas; Kraemer, Roland  
CS Anorganisch-Chemisches Institut, Karl-Rupprechts University of Heidelberg, Heidelberg, 69120, Germany  
SO Bioorganic & Medicinal Chemistry Letters (2003), 13(15), 2489-2492  
CODEN: BMCLE8; ISSN: 0960-894X  
PB Elsevier Science B.V.  
DT Journal  
LA English  
LC CASREACT 139:292469  
AB PNAs with terminal modifications of varying structure and charge were synthesized and their binding to DNA was studied. A variation in thermal stability of 19.8° C has been observed between the least and the most stable PNA-DNA duplexes. The most stable duplex melts

7.7° C higher than the duplex of the corresponding non-modified PNA and complementary DNA. It has been shown that sequence fidelity of the PNA conjugate having the highest DNA affinity is significantly better than that of non-modified PNA. The results obtained can be used for the design of PNA probes, whose binding to DNA is sequence independent.

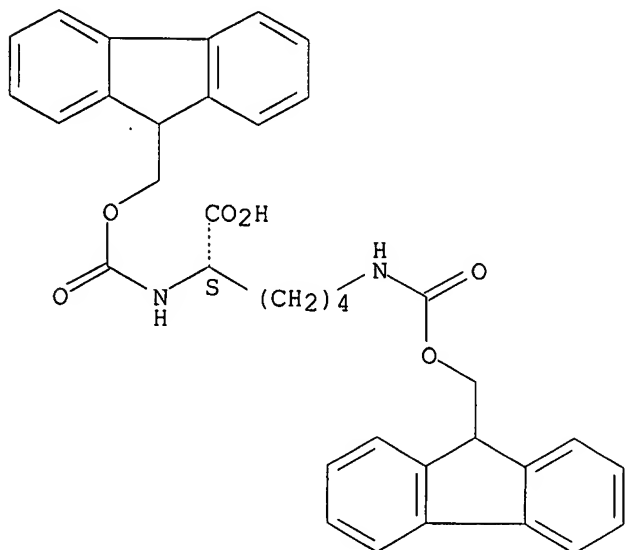
TT 78081-87-5 146982-27-6

RL: RCT (Reactant); RACT (Reactant or reagent)  
(synthesis and DNA binding properties of terminally modified peptide nucleic acids)

RN 78081-87-5 CAPLUS

CN L-Lysine, N2,N6-bis[(9H-fluoren-9-ylmethoxy)carbonyl]- (CA INDEX NAME)

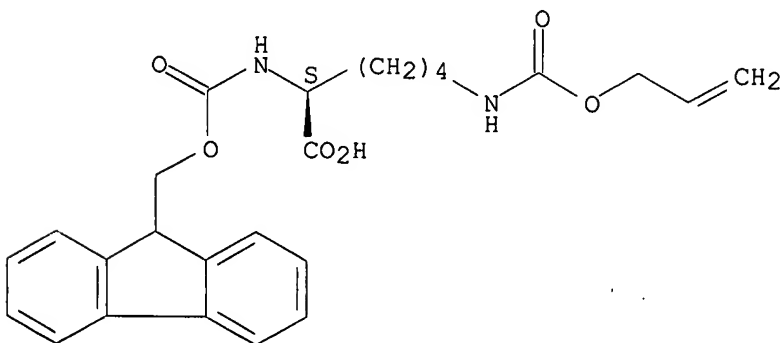
Absolute stereochemistry.



RN 146982-27-6 CAPLUS

CN L-Lysine, N2-[(9H-fluoren-9-ylmethoxy)carbonyl]-N6-[(2-propen-1-yloxy)carbonyl]- (CA INDEX NAME)

Absolute stereochemistry.



RE.CNT 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 4 OF 7 CAPLUS COPYRIGHT 2007 ACS on STN

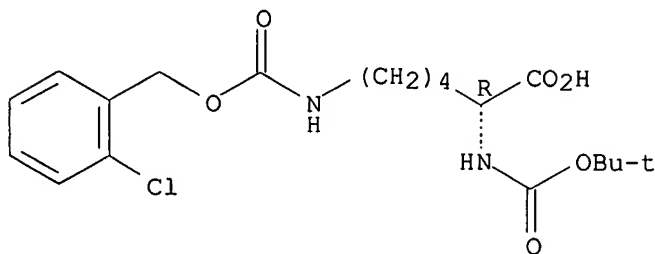
AN 2003:246878 CAPLUS

DN 139:101403

TI Fast, solid-phase synthesis of chiral peptide nucleic acids with a high optical purity by a submonomeric strategy

AU Sforza, Stefano; Tedeschi, Tullia; Corradini, Roberto; Ciavardelli,  
 Domenico; Dossena, Arnaldo; Marchelli, Rosangela  
 CS Dipartimento di Chimica Organica ed Industriale, Universita di Parma,  
 Parma, 43100, Italy  
 SO European Journal of Organic Chemistry (2003), (6), 1056-1063  
 CODEN: EJOCFK; ISSN: 1434-193X  
 PB Wiley-VCH Verlag GmbH & Co. KGaA  
 DT Journal  
 LA English  
 OS CASREACT 139:101403  
 AB The solid-phase synthesis of chiral peptide nucleic acids (PNAs) usually  
 results in partial epimerization of the products, since the  
 $\alpha$ -nitrogen atom of the amino acid is involved in an amidic bond. It  
 is also time-consuming, since all the chiral monomers bearing different  
 nucleobases have to be independently synthesized. In order to prevent  
 racemization and to speed up the synthetic procedure we adopted a  
 submonomeric approach by using a solid-phase, Boc-based (Boc =  
 tert-butoxycarbonyl) PNA synthesis in which the chiral backbone  
 orthogonally  $N\alpha$ -Fmoc-protected (submonomer) (Fmoc  
 = 9-fluorenylmethyloxycarbonyl) was first linked to the growing chain on  
 the resin, followed by Fmoc-deprotection and derivatization with  
 the carboxymethylnucleobase. The submonomer bearing the D-lysine residue  
 was designed by protecting the  $N\alpha$ -(aminoethyl)amino acid moiety with  
 an Fmoc protecting group, compatible with standard Boc chemical, and  
 with the use of an MBHA-PS resin, normally employed for PNA  
 synthesis. Different synthetic pathways towards the desired submonomer  
 were studied by using the amino acid D-lysine as a chiral synthon,  
 obtaining a fast method leading to a high yield and an excellent  
 enantiomeric excess of the submonomer. The solid-phase submonomeric  
 reaction conditions were optimized for the synthesis of a thymineyl  
 PNA dimer and then used to synthesize two different chiral PNAs.  
 In this way two advantages were obtained: a lower degree of racemization  
 in the coupling step during the solid-phase synthesis and the possibility  
 of using the same submonomer for every different nucleobase. All the  
 D-lysine-based chiral PNAs were obtained in good yields and, as compared  
 with PNAs synthesized by other coupling methods, showed the highest  
 optical purity reported so far.  
 IT 57096-11-4  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (asym. solid-phase synthesis of D-lysine-based chiral peptide nucleic  
 acids with by submonomeric strategy)  
 RN 57096-11-4 CAPLUS  
 CN D-Lysine, N6-[[[(2-chlorophenyl)methoxy]carbonyl]-N2-[(1,1-  
 dimethylethoxy)carbonyl]- (CA INDEX NAME)

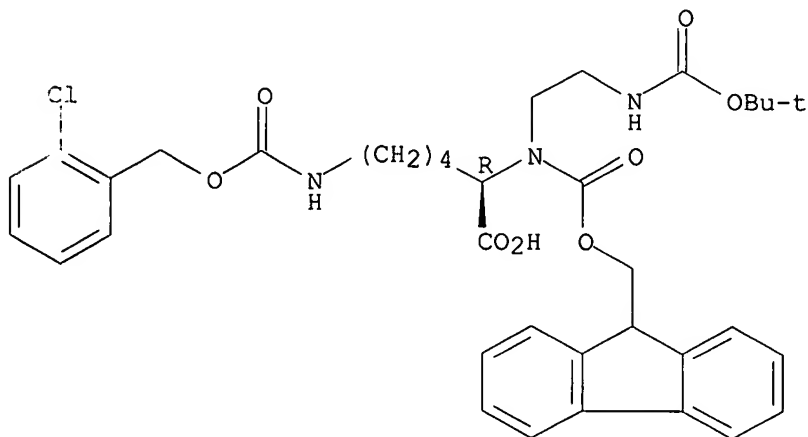
Absolute stereochemistry.



IT 548490-53-5P 548490-54-6P  
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT  
 (Reactant or reagent)  
 (asym. solid-phase synthesis of D-lysine-based chiral peptide nucleic  
 acids with by submonomeric strategy)  
 RN 548490-53-5 CAPLUS

CN 13-Oxa-2,8,11-triazapentadecanoic acid, 7-carboxy-8-[(9H-fluoren-9-ylmethoxy)carbonyl]-14,14-dimethyl-12-oxo-, 1-[(2-chlorophenyl)methyl] ester, (7R)- (9CI) (CA INDEX NAME)

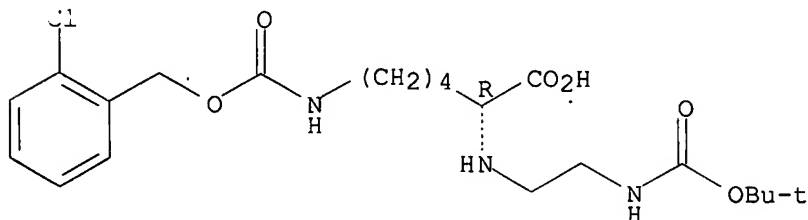
Absolute stereochemistry.



RN 548490-54-6 CAPLUS

CN 13-Oxa-2,8,11-triazapentadecanoic acid, 7-carboxy-14,14-dimethyl-12-oxo-, 1-[(2-chlorophenyl)methyl] ester, (7R)- (CA INDEX NAME)

Absolute stereochemistry.



RE.CNT 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 5 OF 7 CAPLUS COPYRIGHT 2007 ACS on STN

AN 2002:181001 CAPLUS

DN 136:242895

TI Molecular beacons based on peptide nucleic acid probes and their preparation and uses

IN Coull, James M.; Gildea, Brian D.; Hyldig-Nielsen, Jens J.

PA Boston Probes, Inc., USA

SO U.S., 54 pp., Cont.-in-part of U.S. Ser. No. 958,532, abandoned.

CODEN: USXXAM

PT Patent

LA English

FAN.CNT 2

|      | PATENT NO.     | KIND | DATE     | APPLICATION NO. | DATE         |
|------|----------------|------|----------|-----------------|--------------|
| PI   | US 6355421     | B1   | 20020312 | US 1998-179298  | 19981027     |
|      | US 2003036059  | A1   | 20030220 | US 2001-888341  | 20010622 <-- |
|      | US 6528267     | B2   | 20030304 |                 |              |
|      | US 2003232327  | A1   | 20031218 | US 2003-376559  | 20030228 <-- |
|      | US 6949343     | B2   | 20050927 |                 |              |
| PRAI | US 1997-958532 | B2   | 19971027 |                 |              |
|      | US 1998-179298 | A3   | 19981027 |                 |              |

US 2001-888341 A1 20010622

This invention is directed to methods, kits and compns. pertaining to PNA Mol. Beacons. PNA Mol. Beacons comprise self-complementary arm segments and flexible linkages which promote intramol. or intermol. interactions. In the absence of a target sequence, PNA Mol. Beacons facilitate efficient energy transfer between the linked donor and acceptor moieties of the probe. Upon hybridization of the probe to a target sequence, there is a measurable change in at least one property of at least one donor or acceptor moiety of the probe which can be used to detect, identify or quantitate the target sequence in a sample. Synthesis of FRET dye pair-labeled PNAs is described. Optimization expts. for hybridization with PNA mol. probes are described.

IT 105047-45-8P, N- $\alpha$ -(Fmoc)-N- $\epsilon$ -(NH<sub>2</sub>)-L-Lysine-OH 146998-27-8P

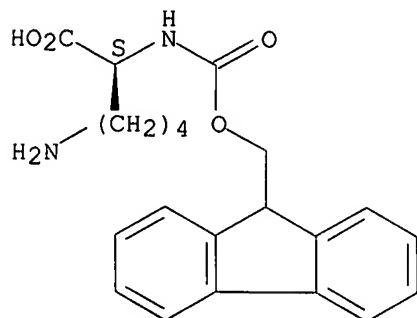
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation and reactions of; mol. beacons based on peptide nucleic acid probes and their preparation and uses)

RN 105047-45-8 CAPLUS

CN L-Lysine, N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (CA INDEX NAME)

Absolute stereochemistry.

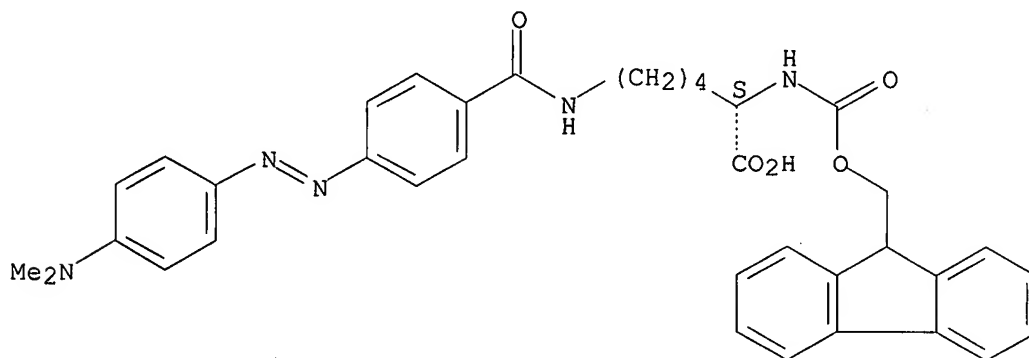


RN 146998-27-8 CAPLUS

CN L-Lysine, N6-[4-[[4-(dimethylamino)phenyl]azo]benzoyl]-N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.



IT 71989-26-9

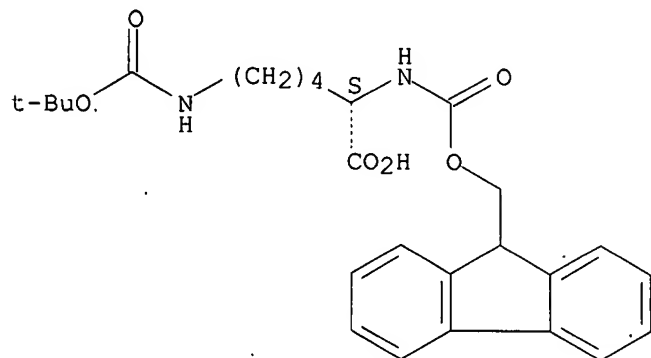
RL: RCT (Reactant); RACT (Reactant or reagent)

(reactions of; mol. beacons based on peptide nucleic acid probes and their preparation and uses)

71989-26-9 CAPLUS

CN L-Lysine, N6-[(1,1-dimethylethoxy)carbonyl]-N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (CA INDEX NAME)

Absolute stereochemistry.

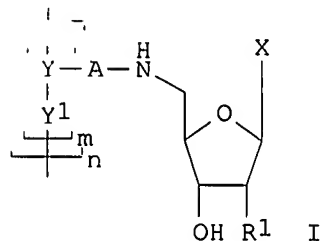


RE.CNT 154 THERE ARE 154 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 6 OF 7 CAPLUS COPYRIGHT 2007 ACS on STN  
AN 2001:923814 CAPLUS  
DN 136:54025  
TI Preparation of peptide nucleoside derivatives as antisense molecules  
IN Inoue, Yoshihisa; Wada, Takehiko  
PA Japan Science and Technology Corporation, Japan  
SO PCT Int. Appl., 77 pp.  
CODEN: PIXXD2  
DT Patent  
LA Japanese  
CAN.CNT 1

|      | PATENT NO.  | KIND | DATE     | APPLICATION NO. | DATE         |
|------|---|------|----------|-----------------|--------------|
| PI   | WO 2001096355   | A1   | 20011220 | WO 2001-JP5011  | 20010613     |
|      | W: JP, US   |      |          |                 |              |
|      | RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR                            |      |          |                 |              |
|      | EP 1295891  | A1   | 20030326 | EP 2001-938631  | 20010613 <-- |
|      | R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR |      |          |                 |              |
|      | US 2003208050   | A1   | 20031106 | US 2002-311048  | 20021212 <-- |
|      | US 6872809  | B2   | 20050329 |                 |              |
| PRAI | JP 2000-177428  | A    | 20000613 |                 |              |
|      | WO 2001-JP5011  | W    | 20010613 |                 |              |

GI



AB Peptide nucleoside derivs. (peptide nucleic acid, PNA)

represented by the following general formula [I; wherein Xs are the same or different and represent pyrimidine, purine nucleic acid base or derivative(s) thereof; Y and Y' are the same or different and represent at least one amino acid or amino acid derivative selected from the group consisting of serine, ornithine, aspartic acid, glutamic acid, lysine, arginine, cysteine,  $\delta$ -hydroxylysine, N-aminoethylglycine, N-aminoethylserine, N-aminoethyllysine, N-aminoethylornithine, N-aminoethylaspartic acid, N-aminoethylglutamic acid, homoglutamic acid,  $\beta$ -thiocarbonylaspartic acid,  $\gamma$ -thiocarbonylglutamic acid and  $\delta$ -thiocarbonylhomoglutamic acid; R1 represents hydrogen or hydroxy; A represents a single bond, carbonyl or thiocarbonyl; m is an integer of from 0 to 5; and n is an integer of from 1 to 100.] are prepared. These compds. exhibit base specific recognition of nucleic acid base sequences with high affinity than natural nucleic acids and are not hydrolyzed easily by enzymes in vivo and useful as antisense mols. for gene therapy of cancer or genetic diseases (no data). They can also irreversibly control the on-off switching of gene expression from anti to syn or syn to anti direction by the influence of pH, light, temperature, or concentration or presence

of alkaline earth or transition metal or sugar. Thus, 0.454 g pentachlorophenyl trichloroacetate and 0.174 mL diisopropylethylamine were added to a solution of 0.129 g poly(L-glutamine) in 20 mL DMF at 0° with stirring, and after 1 h treated with 0.267 g 5'-amino-5'-deoxyuridine, and heated at 60° for 10 h to give 0.314 g poly[Ny-(5'-deoxy-5'-aminouracyl)-L-glutamine].

IT 292616-42-3P 380911-88-6P 380911-92-2P

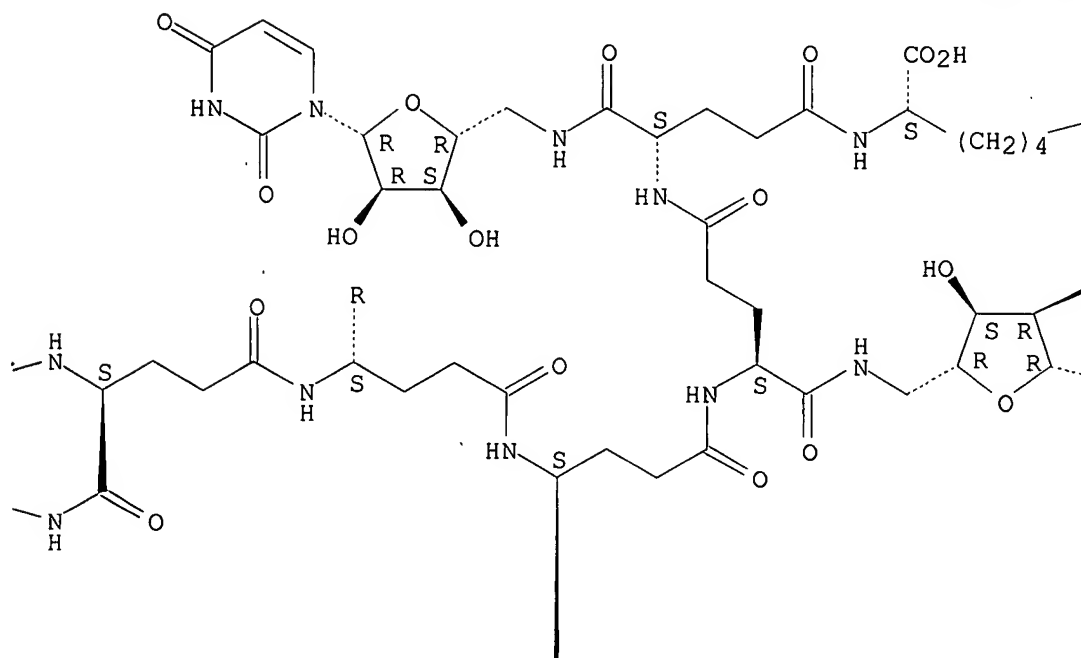
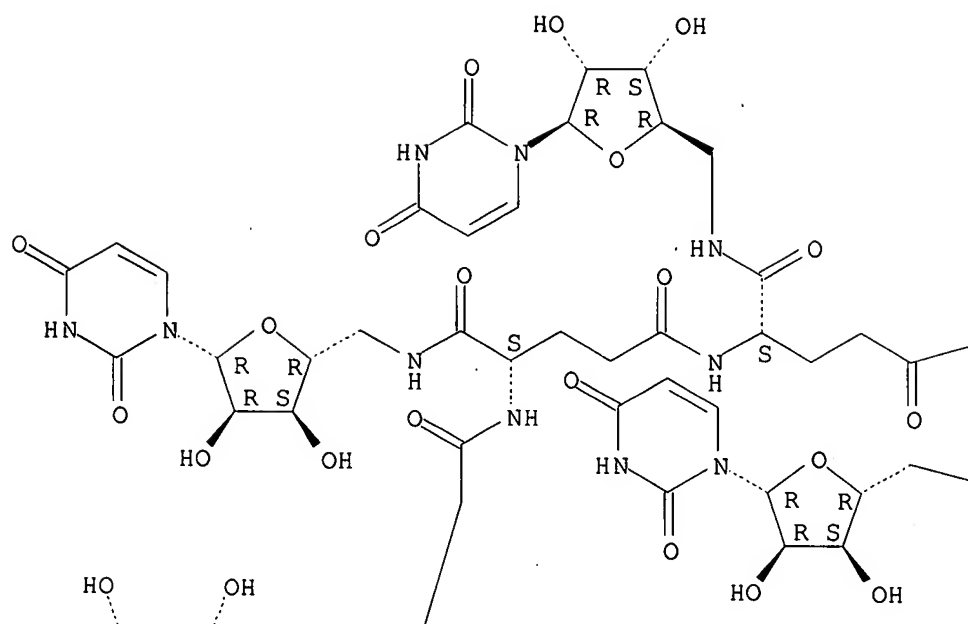
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(preparation of peptide nucleoside derivs. as antisense mols. for gene therapy of cancer or genetic diseases)

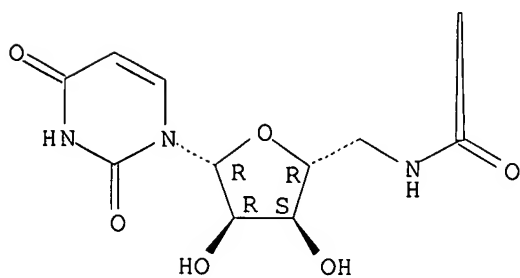
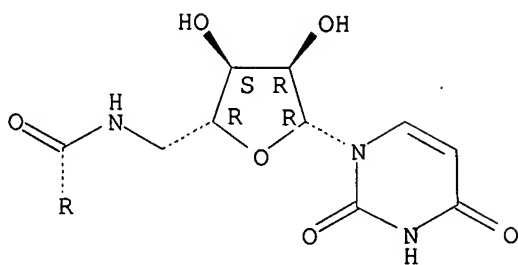
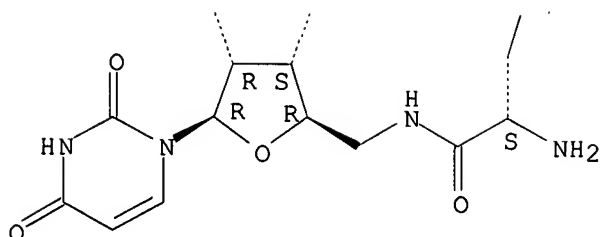
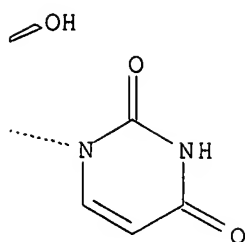
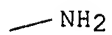
RN 292616-42-3 CAPLUS

CN L-Lysine, N-(5'-deoxyuridin-5'-yl)-L- $\alpha$ -glutaminyL-N-(5'-deoxyuridin-5'-yl)-L- $\alpha$ -glutaminyL-N-(5'-deoxyuridin-5'-yl)-L- $\alpha$ -glutaminyL-N-(5'-deoxyuridin-5'-yl)-L- $\alpha$ -glutaminyL-N-(5'-deoxyuridin-5'-yl)-L- $\alpha$ -glutaminyL-N-(5'-deoxyuridin-5'-yl)-L- $\alpha$ -glutaminyL-, monohydrochloride (9CI) (CA INDEX NAME)

Absolute stereochemistry.







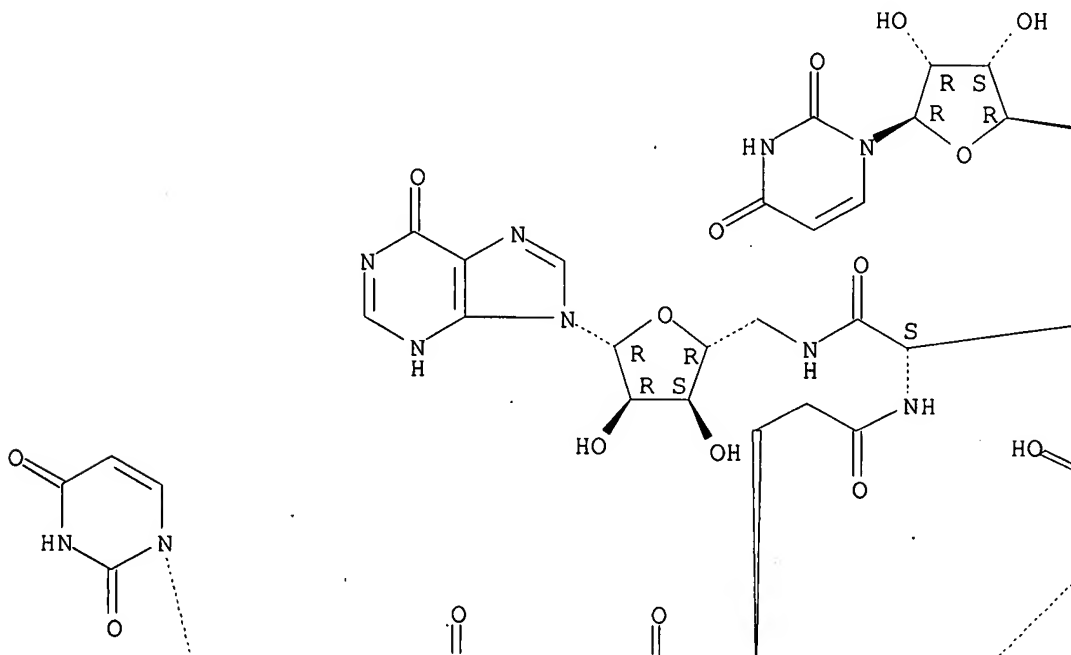
● HCl

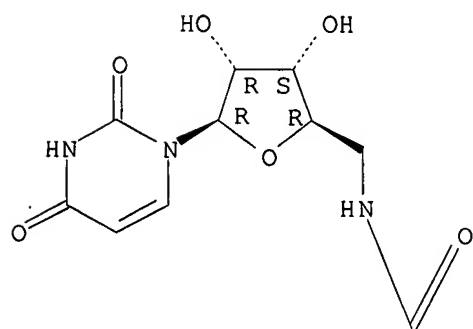
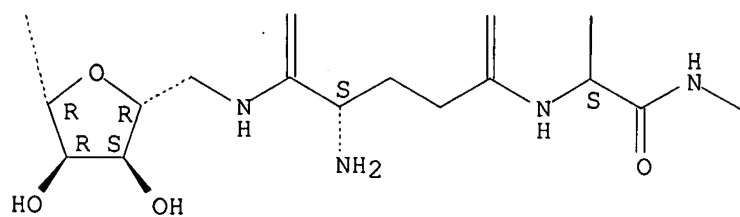
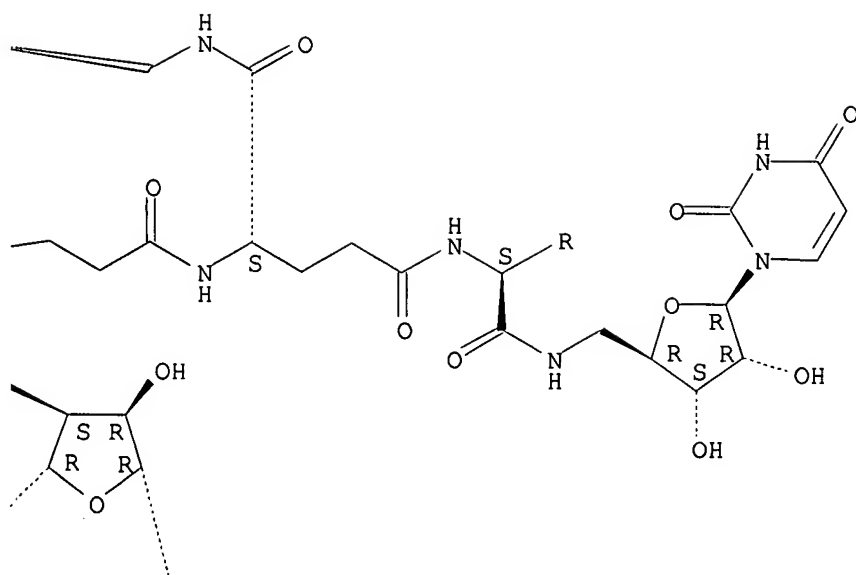
380911-88-6 CAPLUS

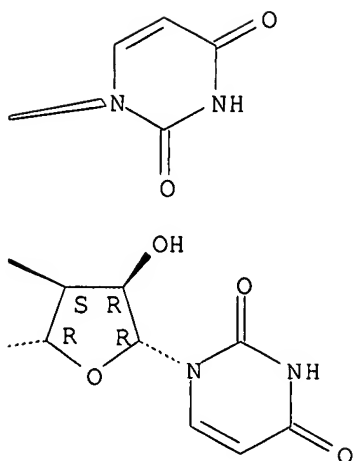
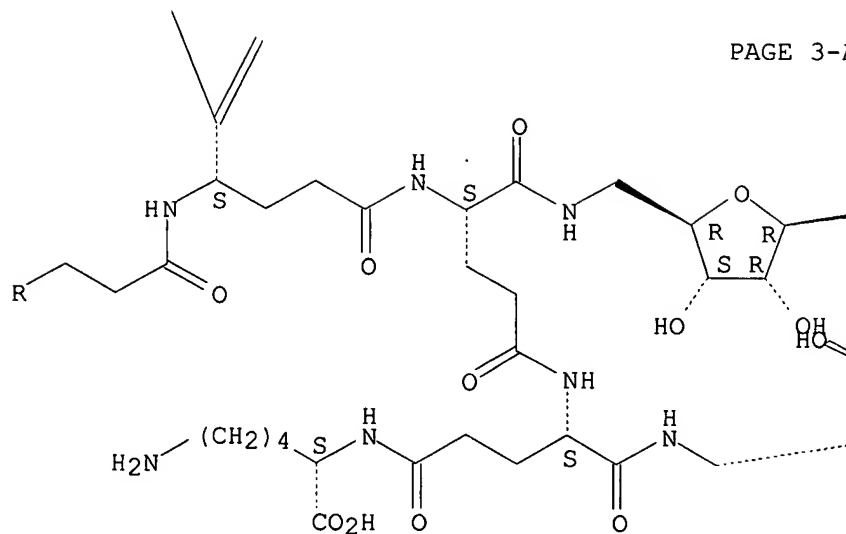
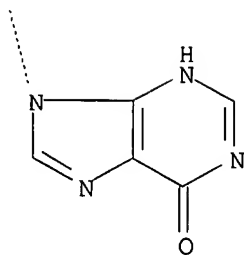
L-Lysine, N-(5'-deoxyuridin-5'-yl)-L- $\alpha$ -glutaminyl-N-(5'-deoxyinosin-5'-yl)-L- $\alpha$ -glutaminyl-N-(5'-deoxyinosin-5'-yl)-L- $\alpha$ -glutaminyl-N-(5'-deoxyuridin-5'-yl)-L- $\alpha$ -glutaminyl-N-(5'-deoxyuridin-5'-yl)-L- $\alpha$ -glutaminyl-N-(5'-deoxyuridin-5'-yl)-L- $\alpha$ -glutaminyl-N-(5'-deoxyuridin-5'-yl)-L- $\alpha$ -glutaminyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



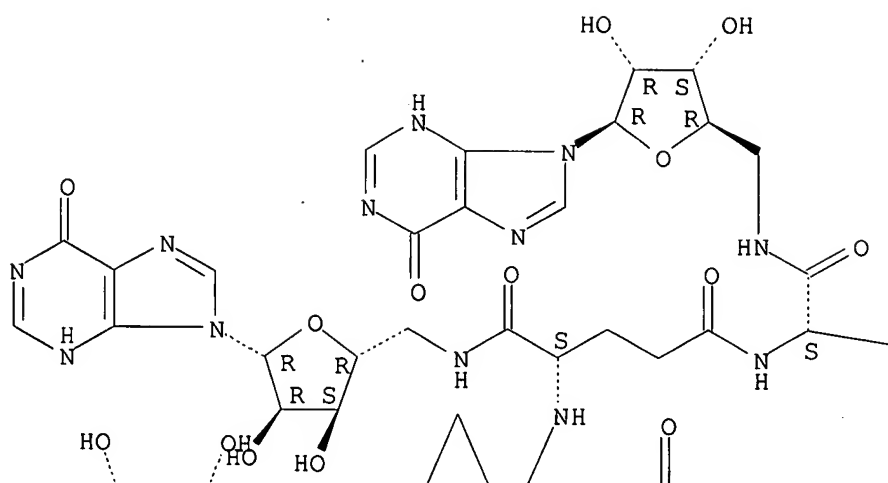


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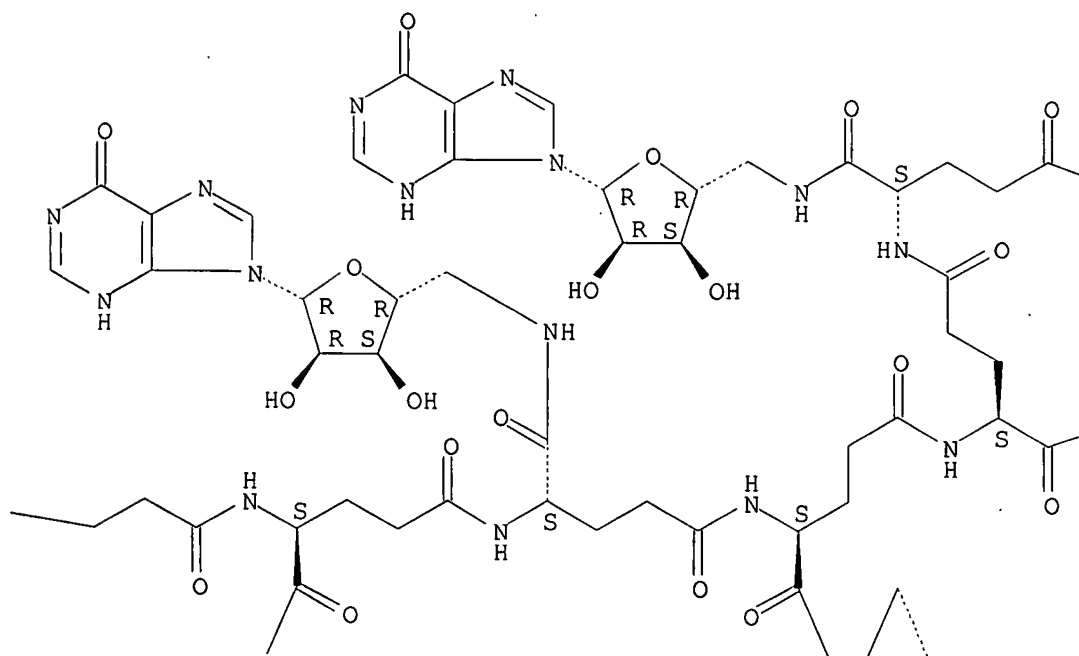
glutaminy- (9CI) (CA INDEX NAME)

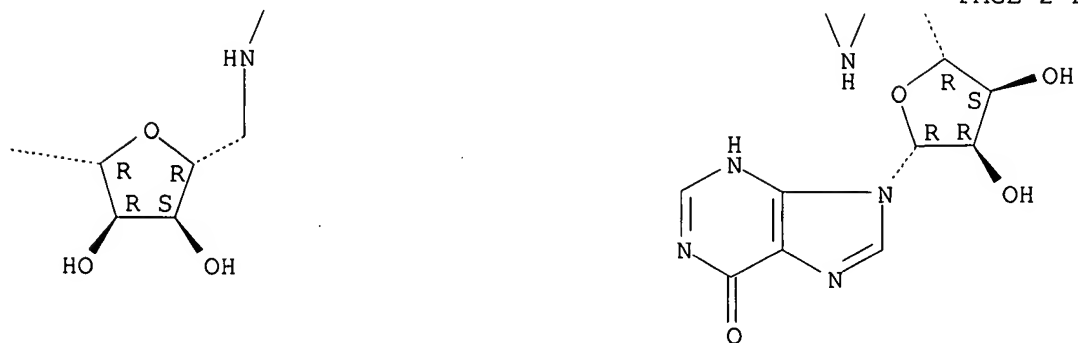
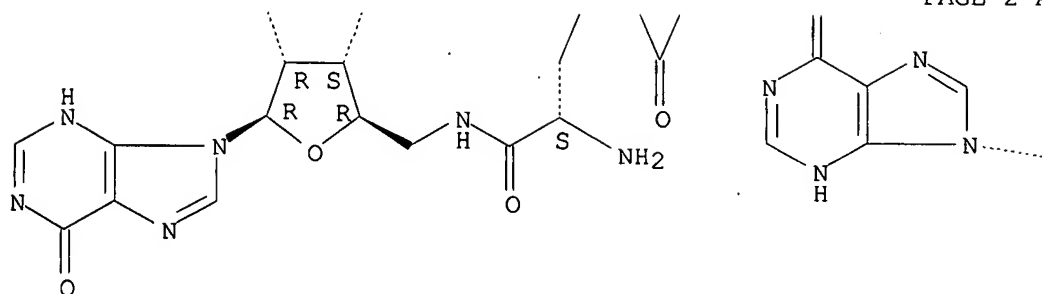
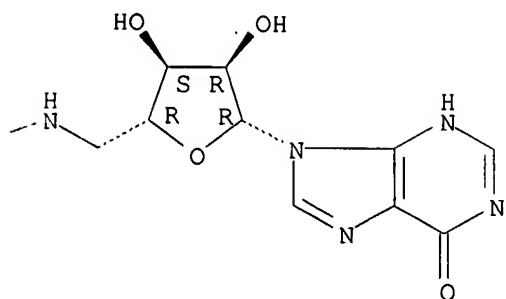
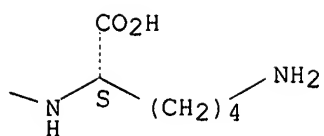
Absolute stereochemistry.

PAGE 1-A



PAGE 1-B





RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

19 ANSWER 7 OF 7 CAPLUS COPYRIGHT 2007 ACS on STN  
AN 1999:626412 CAPLUS  
DN 131:253322  
TI Methods, kits and compositions pertaining to detection complexes for  
nucleic acid targets

IN Coull, James D.; Gildea, Brian D.; Hyldig-Nielsen, Jens J.  
 PA Boston Probes, Inc., USA  
 SO PCT Int. Appl., 123 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 MAN. CNT 1

| PATENT NO.  | KIND | DATE     | APPLICATION NO. | DATE         |
|---|------|----------|-----------------|--------------|
| WO 9949293  | A2   | 19990930 | WO 1999-US6422  | 19990324     |
| WO 9949293  | A3   | 20000406 |                 |              |
| W: AL, AU, BA, BB, BG, BR, CA, CN, CU, CZ, EE, GE, HU, IL, IS, JP, KP, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, TJ, TR, UA, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM |      |          |                 |              |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG                            |      |          |                 |              |
| AU 9930125  | A    | 19991018 | AU 1999-30125   | 19990324     |
| EP 1064399  | A2   | 20010103 | EP 1999-911496  | 19990324     |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI   |      |          |                 |              |
| JP 2002507434   | T    | 20020312 | JP 2000-538214  | 19990324     |
| US 6361942  | B1   | 20020326 | US 1999-275848  | 19990324     |
| US 6607889  | B1   | 20030819 | US 2001-867345  | 20010529 <-- |
| PRAI US 1998-79211P   | P    | 19980324 |                 |              |
| US 1999-275848  | A1   | 19990324 |                 |              |
| WO 1999-US6422  | W    | 19990324 |                 |              |

This invention is directed to methods, kits and compns. which utilize Detection Complexes to detect or identify the presence, absence or quantity of a target mol. in sample of interest. A Detection Complex comprises at least two component polymers and at least one set of donor and acceptor moieties. To each of at least two component polymers is linked at least one moiety of a set of donor and acceptor moieties, such that formation of the complex facilitates transfer of energy between donor and acceptor moieties of each set in a manner which, in an assay, produces changes in detectable signal which can be correlated with the presence/absence of quantity of target sequence and/or target mol. of interest in the sample. The Detection Complexes and PCR detection Complexes of this invention are primarily designed to dissociate as a direct or indirect consequence of the hybridization of one or more segments of a component polymer to a target sequence of a target mol. Because the component polymers of a Detection Complex will preferably dissociate, the attached donor and acceptor moieties, which are independently attached to different polymers, can become far more separated in space as compared with unimol. Beacon probes such as Mol. Beacons or Linear Beacons. As a consequence, the efficiency of energy transfer will be far more substantially altered as compared with unimol. probes wherein the donor and acceptor moieties are linked to the same polymer and therefore cannot be infinitely separated in space. Thus, the Detection Complexes and PCR Detection Complexes of this invention possess a substantial comparative advantage over unimol. Beacon probes. In still another embodiment, this invention is directed to Substrate Detection Complexes which operate as a substrate for an enzyme to thereby generate changes in detectable signal in a target independent manner. At least one of the component polymers comprises a peptide nucleic acid (PNA), and the donor and acceptor moieties comprise a fluorophore (e.g., fluorescein) and a quencher (e.g., DABCYL), resp., for fluorescence resonance energy transfer.

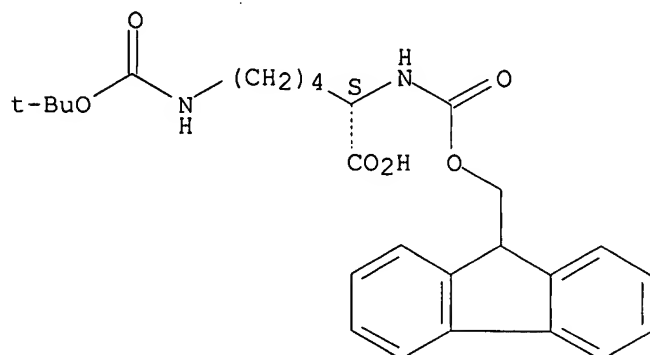
IT 71989-26-9, N- $\alpha$ -(Fmoc)-N- $\epsilon$ -(t-boc)-L-Lysine-OH

RL: RCT (Reactant); RACT (Reactant or reagent)  
 (methods, kits and compns. pertaining to detection complexes for nucleic acid targets)

RN 71989-26-9 CAPLUS

CN L-Lysine, N6-[(1,1-dimethylethoxy)carbonyl]-N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (CA INDEX NAME)

Absolute stereochemistry.



IT 105047-45-8P, N- $\alpha$ -(Fmoc)-N- $\epsilon$ -(NH2)-L-

Lysine-OH 146998-27-8P, Fmoc-K(dabcyl)-OH

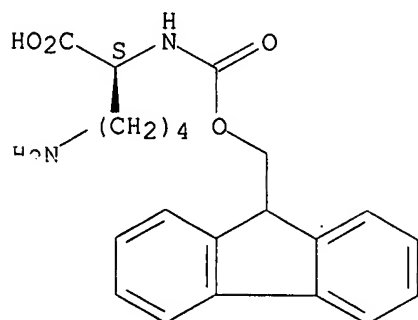
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(methods, kits and compns. pertaining to detection complexes for nucleic acid targets)

RN 105047-45-8 CAPLUS

CN L-Lysine, N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (CA INDEX NAME)

Absolute stereochemistry.



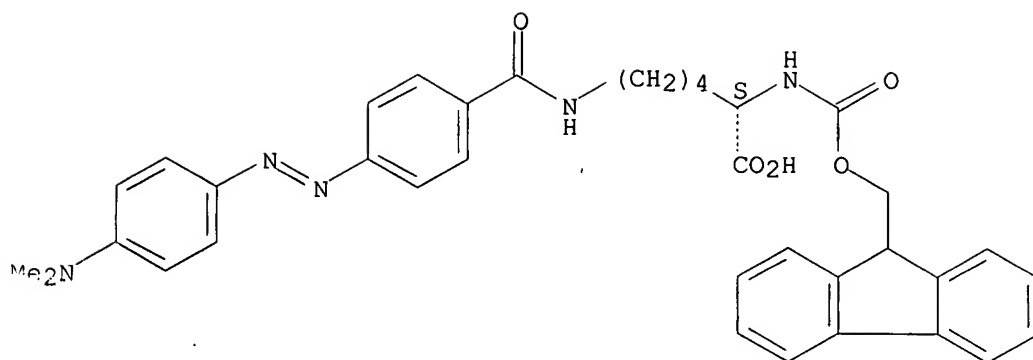
RN 146998-27-8 CAPLUS

CN L-Lysine, N6-[4-[[4-(dimethylamino)phenyl]azo]benzoyl]-N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.





=>

=> d his

(FILE 'HOME' ENTERED AT 12:16:11 ON 10 OCT 2007)

FILE 'REGISTRY' ENTERED AT 12:16:40 ON 10 OCT 2007

L1 STRUCTURE UPLOADED

L2 83880 S L1 FULL

FILE 'CAPLUS' ENTERED AT 12:17:10 ON 10 OCT 2007

L3 107468 S L2

L4 1286 S L3 AND FMOC

L5 27 S L4 AND PNA

L6 27 DUP REM L5 (0 DUPLICATES REMOVED)

L7 27 S L6

L8 7 S L6 AND 2003/PY

=> s 18 and (alloc? or boc?)

20834 ALLOC?

19498 BOC?

L9 3 L8 AND (ALLOC? OR BOC?)

=> d 19 bib abs hitstr 1-3

L9 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN

AN 2003:679388 CAPLUS

DN 139:381726

TI Modulation of the Pharmacokinetic Properties of PNA: Preparation  
of Galactosyl, Mannosyl, Fucosyl, N-Acetylgalactosaminyl, and  
N-Acetylglucosaminyl Derivatives of Aminoethylglycine Peptide Nucleic Acid  
Monomers and Their Incorporation into PNA Oligomers  
AU Hamzavi, Ramin; Dolle, Frederic; Tavitian, Bertrand; Dahl, Otto; Nielsen,  
Peter E.

CC Center for Biomolecular Recognition, Department of Medical Biochemistry  
and Genetics, University of Copenhagen, Copenhagen, DK-2200, Den.

SO Bioconjugate Chemistry (2003), 14(5), 941-954

CODEN: BCCHES; ISSN: 1043-1802

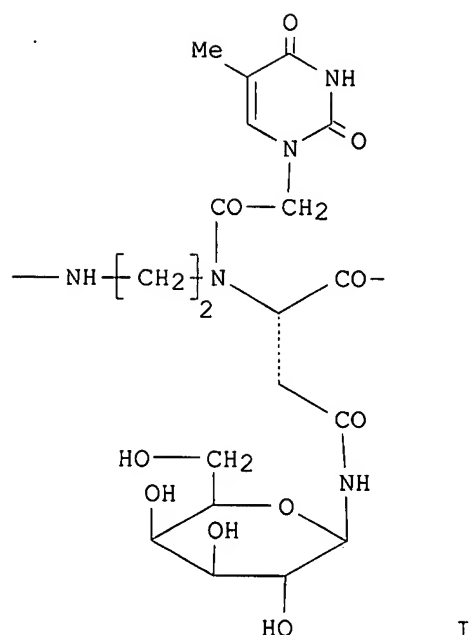
PB American Chemical Society

DT Journal

LA English

OS CASREACT 139:381726

GI



AB A series of N-(2-aminoethyl)- $\alpha$ -amino acid thymine peptide nucleic acid (PNA) monomers bearing glycosylated side chains in the  $\alpha$ -amino acid position (e.g, I) have been synthesized. These include PNA monomers where glycine has been replaced by serine and threonine (O-glycosylated), derivs. of lysine and nor-alanine (C-glycosylated), and amide derivs. of aspartic acid (N-glycosylated). The Boc and Fmoc derivs. of these monomers were used for incorporation in PNA oligomers. Twelve PNA decamers containing the glycosylated units in one, two, or three positions were prepared, and the thermal stability ( $T_m$ ) of their complexes with a complementary RNA was determined. Incorporation of the glycosyl monomers reduced the duplex stability by 0-6° C per substitution. A cysteine was attached to the amino terminus of eight of the PNA decamers (Cys-CTCATACTCT-NH<sub>2</sub>) for easy conjugation to a [18F]radiolabeled N-(4-fluorobenzyl)-2-bromoacetamide. The in vivo biodistribution of these PNA oligomers was determined in rat 2 h after i.v. administration. Most of the radioactivity was recovered in the kidneys and in the urine. However, N-acetylgalactosamine (and to a lesser extent galactose and mannose)-modified PNAs were effectively targeting the liver (40-fold over unmodified PNA). Thus, the pharmacodistribution in rats of PNA oligomers can be profoundly changed by glycosylation. These results could be of great significance for PNA drug development, as they should allow modulation and fine-tuning of the pharmacokinetic profile of a drug lead.

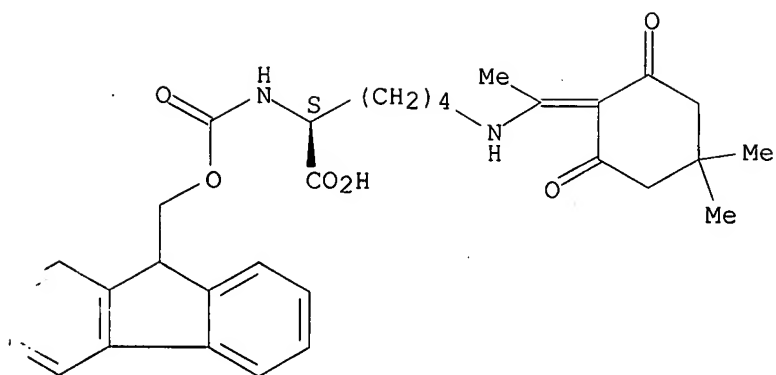
IT 150629-67-7

RL: RCT (Reactant); RACT (Reactant or reagent)  
(preparation of glycosylated monomers for PNA synthesis and their effect on PNA/RNA hybridization or PNA biodistribution)

RN 150629-67-7 CAPLUS

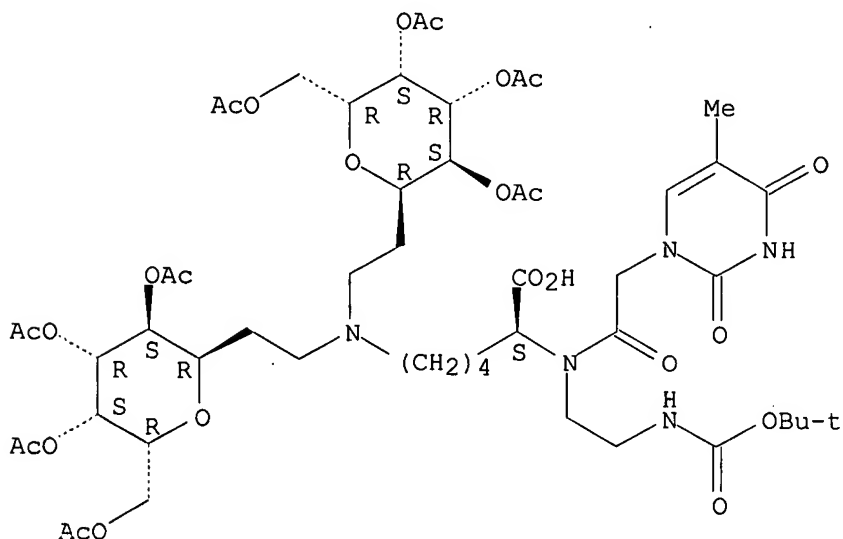
CN L-Lysine, N6-[1-(4,4-dimethyl-2,6-dioxocyclohexylidene)ethyl]-N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (CA INDEX NAME)

Absolute stereochemistry.



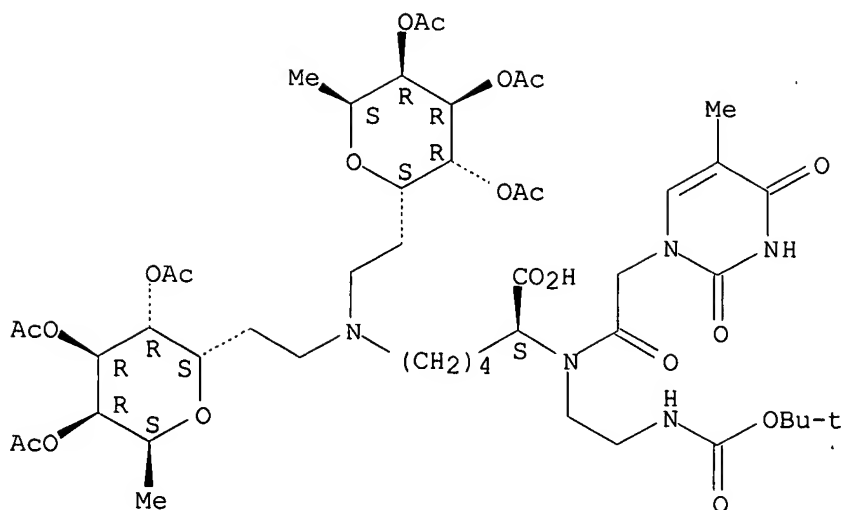
IT 612491-20-0P 612491-21-1P 612491-22-2P  
 612491-23-3P 612491-24-4P 612491-25-5P  
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT  
 (Reactant or reagent)  
 (preparation of glycosylated monomers for PNA synthesis and their  
 effect on PNA/RNA hybridization or PNA  
 biodistribution)  
 RN 612491-20-0 CAPLUS  
 CN L-Lysine, N2-[(3,4-dihydro-5-methyl-2,4-dioxo-1(2H)-pyrimidinyl)acetyl]-N2-  
 [2-[[[(1,1-dimethylethoxy)carbonyl]amino]ethyl]-N6,N6-bis(1,3,4,5-tetra-O-  
 acetyl-2,6-anhydro-7,8-dideoxy-D-glycero-L-galacto-octitol-8-yl)]- (9CI)  
 (CA INDEX NAME)

Absolute stereochemistry.



RN 612491-21-1 CAPLUS  
 CN L-Lysine, N2-[(3,4-dihydro-5-methyl-2,4-dioxo-1(2H)-pyrimidinyl)acetyl]-N2-  
 [2-[[[(1,1-dimethylethoxy)carbonyl]amino]ethyl]-N6,N6-bis(3,4,5-tri-O-  
 acetyl-2,6-anhydro-1,7,8-trideoxy-L-glycero-D-galacto-octitol-8-yl)]- (9CI)  
 (CA INDEX NAME)

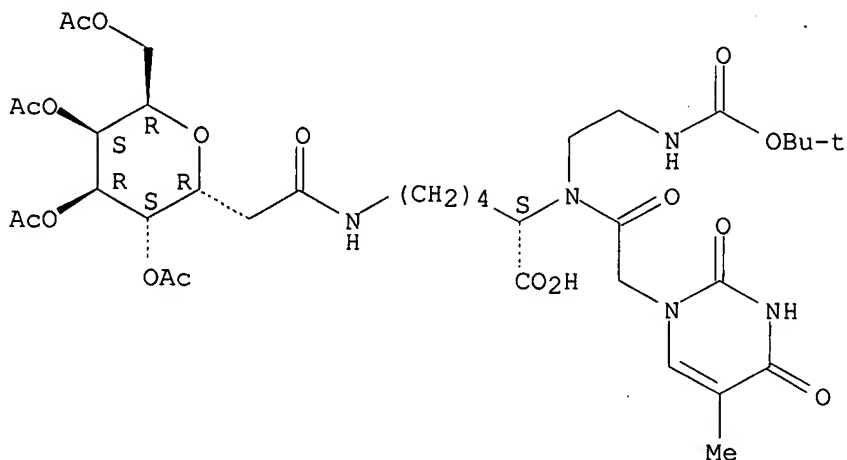
Absolute stereochemistry.



612491-22-2 CAPLUS

L-Lysine, N2-[(3,4-dihydro-5-methyl-2,4-dioxo-1(2H)-pyrimidinyl)acetyl]-N2-[2-[[[(1,1-dimethylethoxy)carbonyl]amino]ethyl]-N6-(4,5,6,8-tetra-O-acetyl-3,7-anhydro-2-deoxy-D-glycero-L-glucopyranosyl)-(9CI) (CA INDEX NAME)

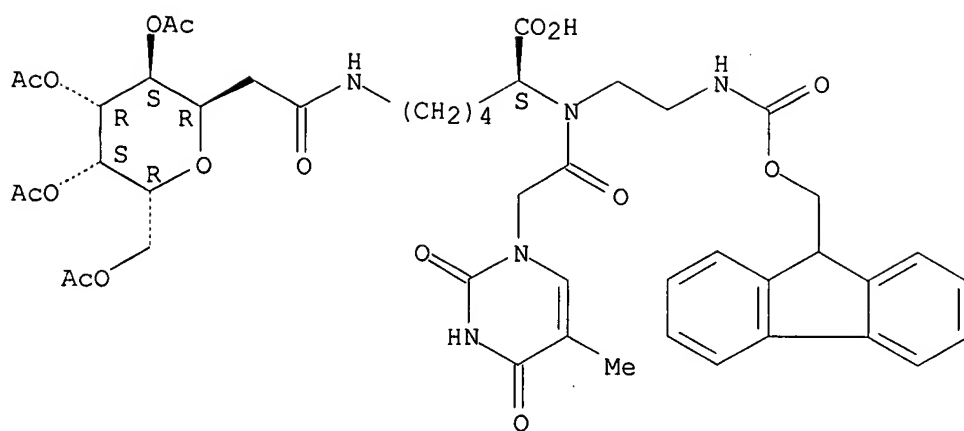
Absolute stereochemistry.



612491-23-3 CAPLUS

L-Lysine, N2-[(3,4-dihydro-5-methyl-2,4-dioxo-1(2H)-pyrimidinyl)acetyl]-N2-[2-[[[(9H-fluoren-9-ylmethoxy)carbonyl]amino]ethyl]-N6-(4,5,6,8-tetra-O-acetyl-3,7-anhydro-2-deoxy-D-glycero-L-glucopyranosyl)-(9CI) (CA INDEX NAME)

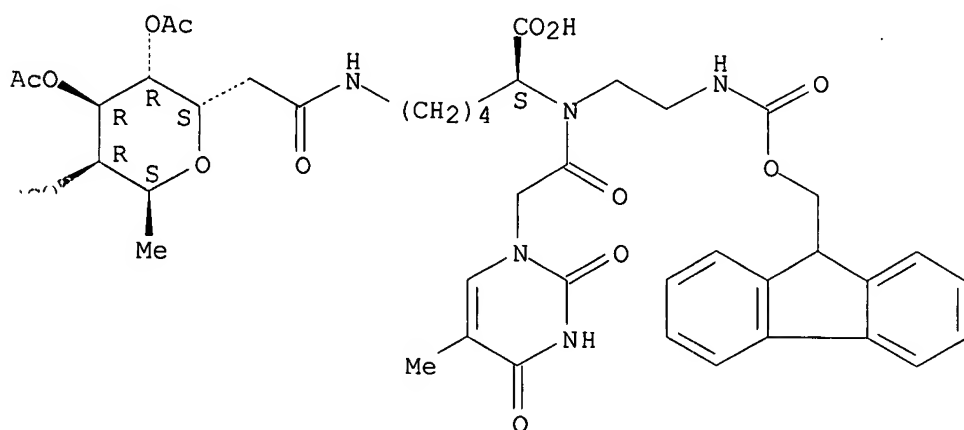
Absolute stereochemistry.



RN 612491-24-4 CAPLUS

CN L-Lysine, N2-[(3,4-dihydro-5-methyl-2,4-dioxo-1(2H)-pyrimidinyl)acetyl]-N2-[2-[[[(9H-fluoren-9-ylmethoxy)carbonyl]amino]ethyl]-N6-(4,5,6-tri-O-acetyl-3,7-anhydro-2,8-dideoxy-L-glycero-D-gluco-octonoyl)-(9CI) (CA INDEX NAME)

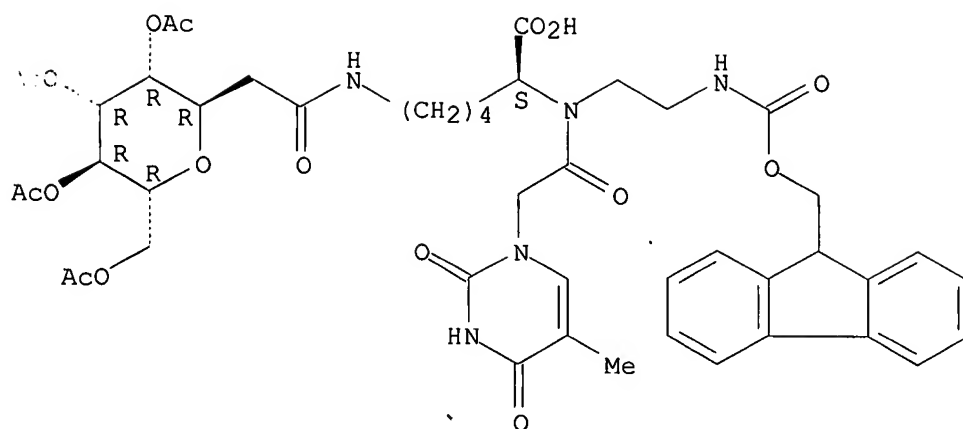
Absolute stereochemistry.



RN 612491-25-5 CAPLUS

CN L-Lysine, N2-[(3,4-dihydro-5-methyl-2,4-dioxo-1(2H)-pyrimidinyl)acetyl]-N2-[2-[[[(9H-fluoren-9-ylmethoxy)carbonyl]amino]ethyl]-N6-(4,5,6,8-tetra-O-acetyl-3,7-anhydro-2-deoxy-D-glycero-D-talo-octonoyl)-(9CI) (CA INDEX NAME)

Absolute stereochemistry.



RE.CNT 40 THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

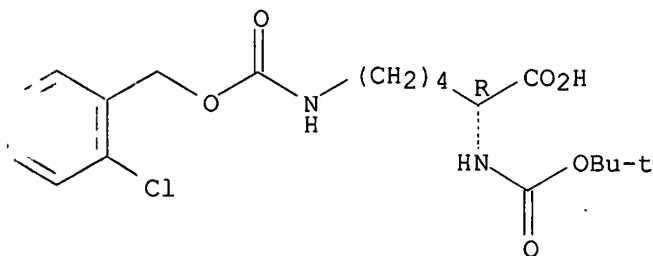
1.9 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN  
 1.1 2003:246878 CAPLUS  
 1.2 139:101403  
 1.1 Fast, solid-phase synthesis of chiral peptide nucleic acids with a high optical purity by a submonomeric strategy  
 AU Sforza, Stefano; Tedeschi, Tullia; Corradini, Roberto; Ciavardelli, Domenico; Dossena, Arnaldo; Marchelli, Rosangela  
 CS Dipartimento di Chimica Organica ed Industriale, Universita di Parma, Parma, 43100, Italy  
 SO European Journal of Organic Chemistry (2003), (6), 1056-1063  
 CODEN: EJOCFK; ISSN: 1434-193X  
 PB Wiley-VCH Verlag GmbH & Co. KGaA  
 DT Journal  
 LA English  
 OS CASREACT 139:101403  
 AB The solid-phase synthesis of chiral peptide nucleic acids (PNAs) usually results in partial epimerization of the products, since the  $\alpha$ -nitrogen atom of the amino acid is involved in an amidic bond. It is also time-consuming, since all the chiral monomers bearing different nucleobases have to be independently synthesized. In order to prevent racemization and to speed up the synthetic procedure we adopted a submonomeric approach by using a solid-phase, Boc-based (Boc = tert-butoxycarbonyl) PNA synthesis in which the chiral backbone orthogonally N $\alpha$ -Fmoc-protected (submonomer) (Fmoc = 9-fluorenylmethoxycarbonyl) was first linked to the growing chain on the resin, followed by Fmoc-deprotection and derivatization with the carboxymethylnucleobase. The submonomer bearing the D-lysine residue was designed by protecting the N $\alpha$ -(aminoethyl)amino acid moiety with an Fmoc protecting group, compatible with standard Boc chemical, and with the use of an MBHA-PS resin, normally employed for PNA synthesis. Different synthetic pathways towards the desired submonomer were studied by using the amino acid D-lysine as a chiral synthon, obtaining a fast method leading to a high yield and an excellent enantiomeric excess of the submonomer. The solid-phase submonomeric reaction conditions were optimized for the synthesis of a thymine PNA dimer and then used to synthesize two different chiral PNAs. In this way two advantages were obtained: a lower degree of racemization in the coupling step during the solid-phase synthesis and the possibility of using the same submonomer for every different nucleobase. All the D-lysine-based chiral PNAs were obtained in good yields and, as compared with PNAs synthesized by other coupling methods, showed the highest optical purity reported so far.  
 57096-11-4  
 RL: RCT (Reactant); RACT (Reactant or reagent)

(asym. solid-phase synthesis of D-lysine-based chiral peptide nucleic acids with by submonomeric strategy)

RN 57096-11-4 CAPLUS

CN D-Lysine, N6-[[ (2-chlorophenyl)methoxy]carbonyl]-N2-[(1,1-dimethylethoxy)carbonyl]- (CA INDEX NAME)

Absolute stereochemistry.



IT 548490-53-5P 548490-54-6P

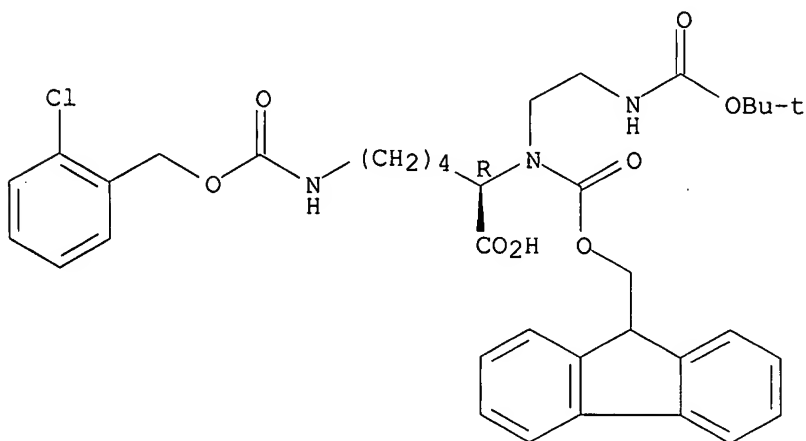
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(asym. solid-phase synthesis of D-lysine-based chiral peptide nucleic acids with by submonomeric strategy)

RN 548490-53-5 CAPLUS

CN 13-Oxa-2,8,11-triazapentadecanoic acid, 7-carboxy-8-[(9H-fluoren-9-ylmethoxy)carbonyl]-14,14-dimethyl-12-oxo-, 1-[(2-chlorophenyl)methyl] ester, (7R)- (9CI) (CA INDEX NAME)

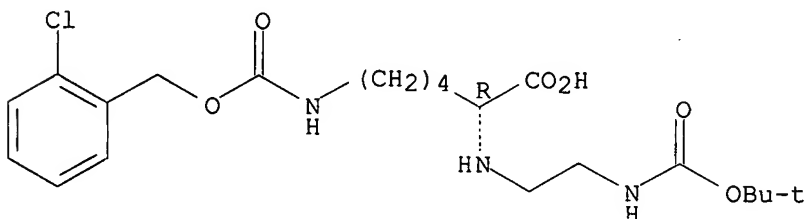
Absolute stereochemistry.



RN 548490-54-6 CAPLUS

CN 13-Oxa-2,8,11-triazapentadecanoic acid, 7-carboxy-14,14-dimethyl-12-oxo-, 1-[(2-chlorophenyl)methyl] ester, (7R)- (CA INDEX NAME)

Absolute stereochemistry.



REF.CNT 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

9 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN  
AN 1999:626412 CAPLUS  
DN 131:253322  
TI Methods, kits and compositions pertaining to detection complexes for  
nucleic acid targets  
IN Coull, James D.; Gildea, Brian D.; Hyldig-Nielsen, Jens J.  
PA Boston Probes, Inc., USA  
SO PCT Int. Appl., 123 pp.  
CODEN: PIXXD2  
DT Patent  
LA English  
FAN.CNT 1

|      | PATENT NO.  | KIND | DATE     | APPLICATION NO. | DATE         |
|------|---|------|----------|-----------------|--------------|
| PI   | WO 9949293  | A2   | 19990930 | WO 1999-US6422  | 19990324     |
|      | WO 9949293  | A3   | 20000406 |                 |              |
|      | W: AL, AU, BA, BB, BG, BR, CA, CN, CU, CZ, EE, GE, HU, IL, IS, JP, KP, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, TJ, TR, UA, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM |      |          |                 |              |
|      | RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG                            |      |          |                 |              |
|      | AU 9930125  | A    | 19991018 | AU 1999-30125   | 19990324     |
|      | EP 1064399  | A2   | 20010103 | EP 1999-911496  | 19990324     |
|      | R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI   |      |          |                 |              |
|      | JP 2002507434   | T    | 20020312 | JP 2000-538214  | 19990324     |
|      | US 6361942  | B1   | 20020326 | US 1999-275848  | 19990324     |
|      | US 6607889  | B1   | 20030819 | US 2001-867345  | 20010529 <-- |
| PRAI | US 1998-79211P  | P    | 19980324 |                 |              |
|      | US 1999-275848  | A1   | 19990324 |                 |              |
|      | WO 1999-US6422  | W    | 19990324 |                 |              |

AB This invention is directed to methods, kits and compns. which utilize Detection Complexes to detect or identify the presence, absence or quantity of a target mol. in sample of interest. A Detection Complex comprises at least two component polymers and at least one set of donor and acceptor moieties. To each of at least two component polymers is linked at least one moiety of a set of donor and acceptor moieties, such that formation of the complex facilitates transfer of energy between donor and acceptor moieties of each set in a manner which, in an assay, produces changes in detectable signal which can be correlated with the presence/absence of quantity of target sequence and/or target mol. of interest in the sample. The Detection Complexes and PCR detection Complexes of this invention are primarily designed to dissociate as a direct or indirect consequence of the hybridization of one or more segments of a component polymer to a target sequence of a target mol. Because the component polymers of a Detection Complex will preferably dissociate, the attached donor and acceptor moieties, which are independently attached to different polymers, can become far more separated in space as compared with unimol. Beacon probes such as Mol. Beacons or Linear Beacons. As a consequence, the efficiency of energy transfer will be far more substantially altered as compared with unimol. probes wherein the donor and acceptor moieties are linked to the same polymer and therefore cannot be infinitely separated in space. Thus, the Detection Complexes and PCR Detection Complexes of this invention possess a substantial comparative advantage over unimol. Beacon probes. In still another embodiment, this invention is directed to Substrate Detection Complexes which operate as a substrate for an enzyme to thereby generate changes in detectable signal in a target independent manner. At least one of the component polymers comprises a peptide nucleic acid (PNA), and the donor and



acceptor moieties comprise a fluorophore (e.g., fluorescein) and a quencher (e.g., DABCYL), resp., for fluorescence resonance energy transfer.

71989-26-9, N- $\alpha$ -(Fmoc)-N- $\epsilon$ -(t-boc)-L-Lysine-OH

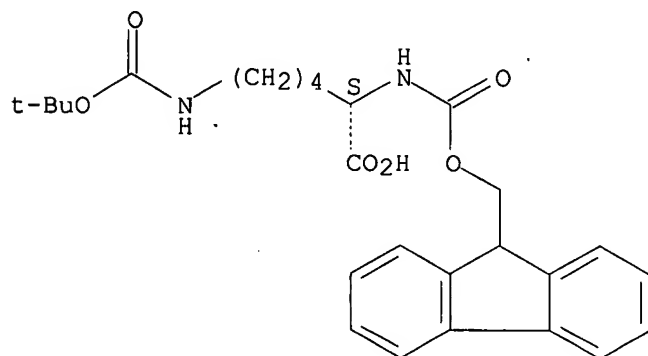
RL: RCT (Reactant); RACT (Reactant or reagent)

(methods, kits and compns. pertaining to detection complexes for nucleic acid targets)

RN 71989-26-9 CAPLUS

CN L-Lysine, N6-[(1,1-dimethylethoxy)carbonyl]-N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (CA INDEX NAME)

Absolute stereochemistry.



IT 105047-45-8P, N- $\alpha$ -(Fmoc)-N- $\epsilon$ -(NH<sub>2</sub>)-L-

Lysine-OH 146998-27-8P, Fmoc-K(dabcyl)-OH

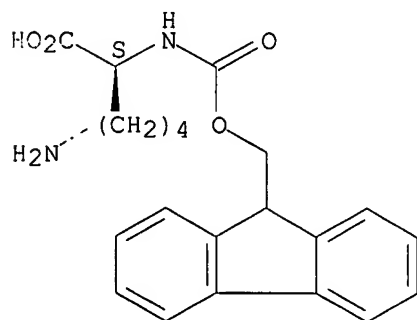
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(methods, kits and compns. pertaining to detection complexes for nucleic acid targets)

RN 105047-45-8 CAPLUS

CN L-Lysine, N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (CA INDEX NAME)

Absolute stereochemistry.

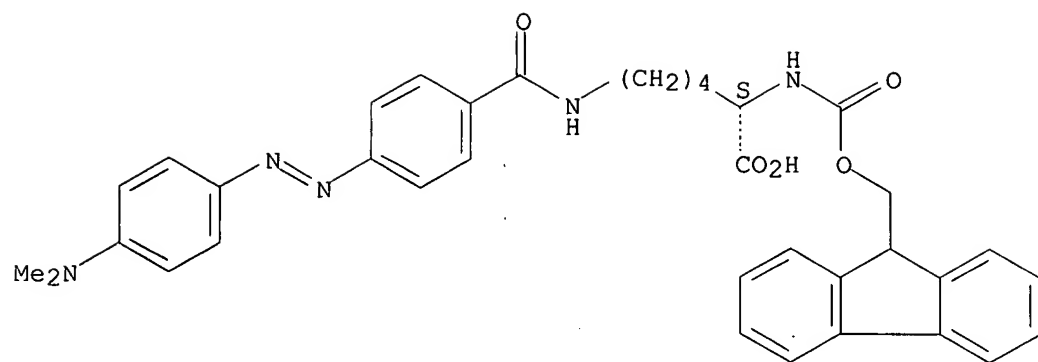


RN 146998-27-8 CAPLUS

CN L-Lysine, N6-[4-[[4-(dimethylamino)phenyl]azo]benzoyl]-N2-[(9H-fluoren-9-ylmethoxy)carbonyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry unknown.



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(FILE 'HOME' ENTERED AT 13:14:04 ON 10 OCT 2007)

FILE 'BIOSIS, MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 13:14:25 ON  
10 OCT 2007

|    |                                      |
|----|--------------------------------------|
| L1 | 3621 S SYNTHESI? (3A) PNA            |
| L2 | 127 S L1 AND LYSINE (4A) FMOC        |
| L3 | 94 S L2 AND (BOC? OR ALLOC?)         |
| L4 | 87 S L3 AND CARBOXYLIC               |
| L5 | 87 DUP REM L4 (0 DUPLICATES REMOVED) |
| L6 | 23 S L5 AND 2003/PY                  |
| L7 | 64 S L5 NOT L6                       |
| L8 | 7 S L5 AND FREE (4A) CARBOXYLIC ACID |